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PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

|      |    |        |   |
|------|----|--------|---|
| NEWS | 1  |        | Web Page for STN Seminar Schedule - N. America  |
| NEWS | 2  | DEC 01 | ChemPort single article sales feature unavailable   |
| NEWS | 3  | JAN 06 | The retention policy for unread STNmail messages<br>will change in 2009 for STN-Columbus and STN-Tokyo          |
| NEWS | 4  | JAN 07 | WPIDS, WPINDEX, and WPIX enhanced Japanese Patent<br>Classification Data  |
| NEWS | 5  | FEB 02 | Simultaneous left and right truncation (SLART) added<br>for CERAB, COMPUAB, ELCOM, and SOLIDSTATE               |
| NEWS | 6  | FEB 02 | GENBANK enhanced with SET PLURALS and SET SPELLING  |
| NEWS | 7  | FEB 06 | Patent sequence location (PSL) data added to USGENE   |
| NEWS | 8  | FEB 10 | COMPENDEX reloaded and enhanced   |
| NEWS | 9  | FEB 11 | WTEXTILES reloaded and enhanced   |
| NEWS | 10 | FEB 19 | New patent-examiner citations in 300,000 CA/Caplus<br>patent records provide insights into related prior<br>art |
| NEWS | 11 | FEB 19 | Increase the precision of your patent queries -- use<br>terms from the IPC Thesaurus, Version 2009.01           |
| NEWS | 12 | FEB 23 | Several formats for image display and print options<br>discontinued in USPATFULL and USPAT2                     |
| NEWS | 13 | FEB 23 | MEDLINE now offers more precise author group fields<br>and 2009 MeSH terms                                      |
| NEWS | 14 | FEB 23 | TOXCENTER updates mirror those of MEDLINE - more<br>precise author group fields and 2009 MeSH terms             |
| NEWS | 15 | FEB 23 | Three million new patent records blast AEROSPACE into<br>STN patent clusters                                    |
| NEWS | 16 | FEB 25 | USGENE enhanced with patent family and legal status<br>display data from INPADOCDB                              |
| NEWS | 17 | MAR 06 | INPADOCDB and INPAFAMDB enhanced with new display<br>formats  |
| NEWS | 18 | MAR 11 | EPFULL backfile enhanced with additional full-text<br>applications and grants                                   |
| NEWS | 19 | MAR 11 | ESBIOBASE reloaded and enhanced   |
| NEWS | 20 | MAR 20 | CAS databases on STN enhanced with new super role<br>for nanomaterial substances                                |
| NEWS | 21 | MAR 23 | CA/Caplus enhanced with more than 250,000 patent<br>equivalents from China                                      |
| NEWS | 22 | MAR 30 | IMSPATENTS reloaded and enhanced  |
| NEWS | 23 | APR 03 | CAS coverage of exemplified prophetic substances<br>enhanced  |
| NEWS | 24 | APR 07 | STN is raising the limits on saved answers  |

NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,  
AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS LOGIN Welcome Banner and News Items

NEWS IPC8      For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 11:33:44 ON 17 APR 2009

=> FIL REGISTRY

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|----------------------|------------------|---------------|
| FULL ESTIMATED COST  | 0.22             | 0.22          |

FILE 'REGISTRY' ENTERED AT 11:34:01 ON 17 APR 2009

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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 15 APR 2009 HIGHEST RN 1135193-69-9

DICTIONARY FILE UPDATES: 15 APR 2009 HIGHEST RN 1135193-69-9

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 9, 2009.

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/support/stngen/stndoc/properties.html>

=> E "DMXAA"/CN 25

|     |       |   |
|-----|-------|---|
| E1  | 1     | DMX 400YB40RBK/CN   |
| E2  | 1     | DMX 7R/CN   |
| E3  | 1 --> | DMXAA/CN  |
| E4  | 1     | DMXAA SODIUM SALT/CN  |
| E5  | 1     | DMXAA-DICLOFENAC MIXTURE/CN                                     |
| E6  | 1     | DMXB-A/CN   |
| E7  | 1     | DMY PROTEIN (ORYZIAS CURVINOTUS GENE DMY)/CN                    |
| E8  | 1     | DMZ/CN  |
| E9  | 3     | DN/CN   |
| E10 | 1     | DN (DISPERSANT)/CN  |
| E11 | 1     | DN (HUMAN PAPILLOMAVIRUS 35 GENE L1 253-NUCLEOTIDE FRAGMENT)/CN |
| E12 | 1     | DN (HUMAN PAPILLOMAVIRUS 39 GENE L1 253-NUCLEOTIDE FRAGMENT)/CN |
| E13 | 1     | DN (HUMAN PAPILLOMAVIRUS 44 GENE L1 244-NUCLEOTIDE FRAGMENT)/CN |
| E14 | 1     | DN (HUMAN PAPILLOMAVIRUS 45 GENE L1 256-NUCLEOTIDE FRAGMENT)/CN |
| E15 | 1     | DN (HUMAN PAPILLOMAVIRUS 51 GENE L1 250-NUCLEOTIDE FRAGMENT)/CN |

E16 1 DN (HUMAN PAPILLOMAVIRUS 56 GENE L1 250-NUCLEOTIDE FRAGMENT)/CN  
 E17 1 DN (HUMAN PAPILLOMAVIRUS 59 GENE L1 253-NUCLEOTIDE FRAGMENT)/CN  
 E18 1 DN (HUMAN PAPILLOMAVIRUS 66 GENE L1 250-NUCLEOTIDE FRAGMENT)/CN  
 E19 1 DN (HUMAN PAPILLOMAVIRUS 68 GENE L1 120-NUCLEOTIDE FRAGMENT)/CN  
 E20 1 DN (HUMAN PROTEIN SERINE/THREONINE KINASE GENE PLUS FLANKS)/CN  
 E21 1 DN (PESTICIDE)/CN  
 E22 2 DN 003/CN  
 E23 1 DN 0081/CN  
 E24 1 DN 02/CN  
 E25 1 DN 099/CN

=> S E3

L1 1 DMXAA/CN

=> DIS L1 1 SQIDE

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2009 ACS on STN

RN 117570-53-3 REGISTRY

CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)

OTHER NAMES:

CN 5,6-Dimethyl-9-oxo-9H-xanthen-4-ylacetic acid

CN 5,6-Dimethylxanthenone-4-acetic acid

CN AS 1404

CN DMXAA

CN NSC 640488

CN Vadimezan

MF C17 H14 O4

CI COM

SR CA

LC STN Files: ADISINSIGHT, ADISNEWS, ANABSTR, BEILSTEIN\*, BIOSIS, CA, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CIN, EMBASE, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, PHAR, PROMT, PROUSDDR, RTECS\*, SYNTHLINE, TOXCENTER, USPAT2, USPATFULL

(\*File contains numerically searchable property data)

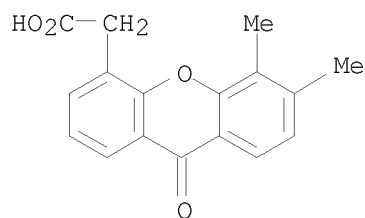
DT.CA CAplus document type: Conference; Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); FORM (Formation, nonpreparative)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

189 REFERENCES IN FILE CA (1907 TO DATE)

4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
189 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> E "GEMCITABINE"/CN 25

|     |   |                                    |
|-----|---|------------------------------------|
| E1  | 1 | GEMCADIOL/CN                       |
| E2  | 1 | GEMCAT 200/CN                      |
| E3  | 1 | --> GEMCITABINE/CN                 |
| E4  | 1 | GEMCITABINE 5'-DIPHOSPHATE/CN      |
| E5  | 1 | GEMCITABINE HYDROCHLORIDE/CN       |
| E6  | 1 | GEMCITABINE TRIPHOSPHATE/CN        |
| E7  | 1 | GEMCO/CN                           |
| E8  | 1 | GEMEDINE/CN                        |
| E9  | 1 | GEMEDIS/CN                         |
| E10 | 1 | GEMEPROST/CN                       |
| E11 | 1 | GEMETREL/CN                        |
| E12 | 1 | GEMEX/CN                           |
| E13 | 1 | GEMEX AGENT 03/CN                  |
| E14 | 1 | GEMFIBROZIL/CN                     |
| E15 | 1 | GEMFIBROZIL 1-O-B-D-GLUCURONIDE/CN |
| E16 | 1 | GEMFIBROZIL GLUCURONIDE/CN         |
| E17 | 1 | GEMFIBROZIL POTASSIUM SALT/CN      |
| E18 | 1 | GEMFIBROZIL SODIUM SALT/CN         |
| E19 | 1 | GEMFIBROZIL-VITAMIN B6 MIXTURE/CN  |
| E20 | 1 | GEMFLEX 1031C/CN                   |
| E21 | 1 | GEMFLEX 307/CN                     |
| E22 | 1 | GEMFLEX 409/CN                     |
| E23 | 1 | GEMGEL 100/CN                      |
| E24 | 1 | GEMGEL 100+/CN                     |
| E25 | 1 | GEMICALCONE A/CN                   |

=> S E3

L2 1 GEMCITABINE/CN

=> DIS L2 1 SQIDE

L2 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2009 ACS on STN

RN 95058-81-4 REGISTRY

CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

OTHER NAMES:

CN 2',2'-Difluoro-2'-deoxycytidine

CN 2',2'-Difluorodeoxycytidine

CN 2'-Deoxy-2',2'-difluorocytidine

CN DDFC

CN DFdC

CN DFdCyd

CN Folfugem

CN Gamcitabine

CN Gemcitabine

CN LY 188011

CN NSC 613327

FS STEREOSEARCH

MF C9 H11 F2 N3 O4

CI COM

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN\*, BIOSIS,  
CA, CAPLUS, CASREACT, CBNB, CHEMCATS, CIN, DDFU, DRUGU, HSDB\*,  
IMSDRUGNEWS, IMSPATENTS, IMSPRODUCT, IMSRESEARCH, IPA, MRCK\*, PATDPASPC,  
PHAR, PROMT, PROUSDDR, PS, RTECS\*, SYNTHLINE, TOXCENTER, USAN, USPAT2,  
USPATFULL

(\*File contains numerically searchable property data)

Other Sources: WHO

DT.CA Caplus document type: Book; Conference; Dissertation; Journal; Patent

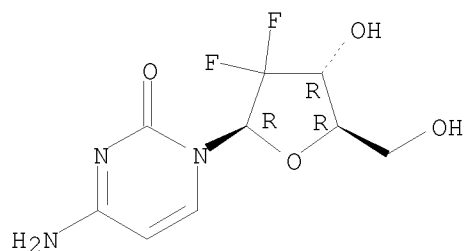
RL.P Roles from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); PRPH (Prophetic); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); PRPH (Prophetic); RACT (Reactant or reagent); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

Absolute stereochemistry. Rotation (+).



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

4554 REFERENCES IN FILE CA (1907 TO DATE)  
 89 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 4594 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> file medline caplus wpids uspatfull

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|----------------------|------------------|---------------|
| FULL ESTIMATED COST  | 15.76            | 15.98         |

FILE 'MEDLINE' ENTERED AT 11:34:54 ON 17 APR 2009

FILE 'CAPLUS' ENTERED AT 11:34:54 ON 17 APR 2009

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FILE 'WPIDS' ENTERED AT 11:34:54 ON 17 APR 2009

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FILE 'USPATFULL' ENTERED AT 11:34:54 ON 17 APR 2009

CA INDEXING COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

=> s 11

L3 370 L1

=> s 12

L4 5732 L2

=> s 13 and 14

L5 13 L3 AND L4

=> d 15 1-13 ibib, abs, hitstr

L5 ANSWER 1 OF 13 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:1250046 CAPLUS

DOCUMENT NUMBER: 149:448110

TITLE: Preparation of Iso CA-4 and analogs as potent cytotoxic agents and inhibitors of polymerization of tubulin

INVENTOR(S): Alami, Mouad; Brion, Jean-Daniel; Provot, Olivier; Peyrat, Jean-Francois; Messaoudi, Samir; Hamze, Abdallah; Giraud, Anne; Bignon, Jerome; Bakala, Joanna; Liu, Jian-Miao

PATENT ASSIGNEE(S): Centre National De La Recherche Scientifique, Fr.

SOURCE: PCT Int. Appl., 78pp.

CODEN: PIXXD2

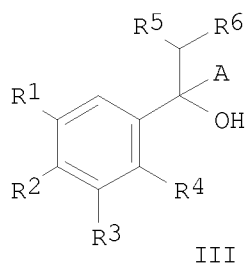
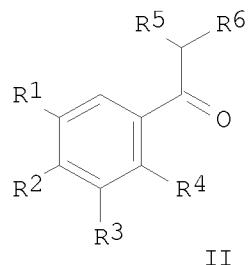
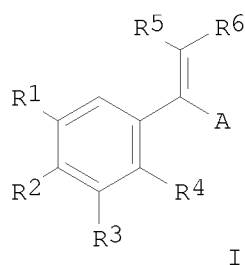
DOCUMENT TYPE: Patent

LANGUAGE: French

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO.             | KIND   | DATE     | APPLICATION NO. | DATE       |
|------------------------|--|----------|-----------------|------------|
| WO 2008122620          | A1   | 20081016 | WO 2008-EP54118 | 20080404   |
| W:                     | AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW |          |                 |            |
| RW:                    | AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM   |          |                 |            |
| FR 2914640             | A1   | 20081010 | FR 2007-54280   | 20070404   |
| PRIORITY APPLN. INFO.: |  |          | FR 2007-54280   | A 20070404 |
| OTHER SOURCE(S):       | MARPAT 149:448110  |          |                 |            |
| GI                     |  |          |                 |            |

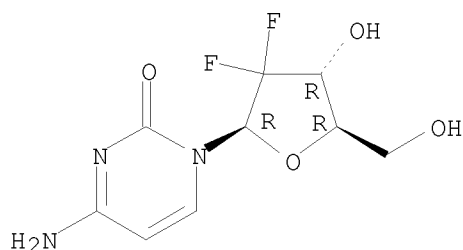


AB Isocombretastatin A-4 and analogs I [R1, R2, R3 = methoxy (possibly substituted by one or more fluorine atoms); R5 = R6 = hydrogen or fluorine; A = ring chosen from (un)substituted aryls and heteroaryl]. The process for the preparation of I comprises: (a) reaction of acetophenone derivative II with an organometallic compound, A-M [M = alkali metal or earth alkaline metal substituted with a halogen]; and (b) reaction of the resulting phenylethanol derivative III with an acid to form I. Thus, Iso-CA-4 [I; A =

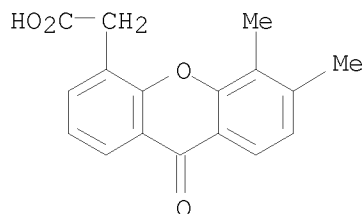
C6H3OH-3-OMe-4, R1 = R2 = R3 = OMe, R4 = R5 = R6 = H (IV)] was prepared from 3,4,5-trimethoxyacetophenone (II; R1 = R2 = R3 = OMe, R4 = R5 = R6 = H) via reaction in PhMe with tert-butyl(5-lithio-2-methoxyphenoxy)dimethylsilane [prepared from tert-butyl(5-iodo-2-methoxyphenoxy)dimethylsilane via lithiation with Me3CLi in hexane], dehydration of III with p-toluenesulfonic acid in CH2Cl2, and desilylation with K2CO3 in MeOH. The cytotoxic activity of IV was determined [IC50 = 2-4 nM vs. HCT116; IC50 = 5 nM vs. K562 cells; IC50 = 2 nM vs. B16F10 cells; IC50 = 8 nM vs. U87 cells; IC50 = 8 nM vs. A549 cells; IC50 = 4.5 nM vs. M435 cells; IC50 = 4 nM vs. M231 cells; IC50 = 2.2  $\mu$ M vs tubulin polymerization].

IT 95058-81-4, Gemcitabine  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (combination chemotherapy antitumor agent; iso CA-4 and analogs as powerful cytotoxic agents and inhibitors of tubulin polymerization)  
 RN 95058-81-4 CAPLUS  
 CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



IT 117570-53-3, DMXAA  
 RL: RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)  
 (reaction of, with iso CA-4 and aminodeoxy-iso-CA-4; iso CA-4 and analogs as powerful cytotoxic agents and inhibitors of tubulin polymerization)  
 RN 117570-53-3 CAPLUS  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



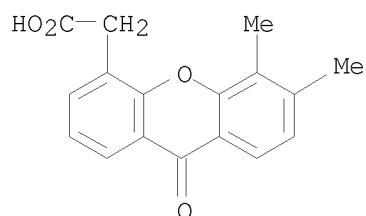
REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 2 OF 13 CAPLUS COPYRIGHT 2009 ACS on STN  
 ACCESSION NUMBER: 2008:473431 CAPLUS  
 DOCUMENT NUMBER: 148:463206  
 TITLE: oncolytic viruses and antiangiogenic agents in the treatment of cancer  
 INVENTOR(S): Karrasch, Matthias; Mescheder, Axel  
 PATENT ASSIGNEE(S): Medigene AG, Germany  
 SOURCE: PCT Int. Appl., 69pp.

CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

| PATENT NO.  | KIND | DATE     | APPLICATION NO. | DATE     |
|---|------|----------|-----------------|----------|
| WO 2008043576   | A1   | 20080417 | WO 2007-EP8930  | 20071015 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW<br>RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM |      |          |                 |          |

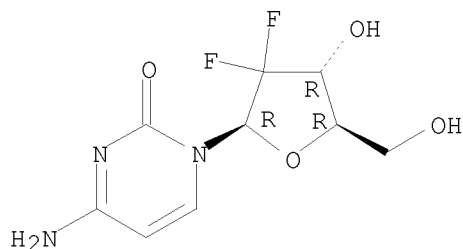
PRIORITY APPLN. INFO.: US 2006-851598P P 20061013  
AB The invention relates to a combination of at least one oncolytic virus and at least one antiangiogenic agent and to the use of this combination in tumor therapy. Intraarterial infusions of oncolytic virus NV1020 to a patient with progressive metastatic colorectal adenocarcinoma followed by CPT-11 plus cetuximab resulted in stabilization of the disease at 6 mo post treatment.  
IT 117570-53-3, DMXAA  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (oncolytic viruses and antiangiogenic agents in treatment of cancer)  
RN 117570-53-3 CAPLUS  
CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



IT 95058-81-4, Gemcitabine  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (oncolytic viruses and antiangiogenic agents in treatment of cancer)  
RN 95058-81-4 CAPLUS  
CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).





REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

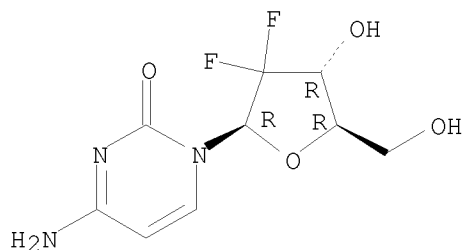
L5 ANSWER 3 OF 13 CAPLUS COPYRIGHT 2009 ACS on STN  
 ACCESSION NUMBER: 2005:984120 CAPLUS  
 DOCUMENT NUMBER: 143:279360  
 TITLE: Methods of detecting CD133 antigen (AC133) expression level and use as biomarker for human cancer diagnosis and therapy monitor  
 INVENTOR(S): Penning, Maarten Tjerk; Van den Broek, Sebastiaan Johannes Jacobus; Voest, Emile Eugene; Beerepoot, Laurens Victor; Mehra, Niven  
 PATENT ASSIGNEE(S): Primagen Holding B. V., Neth.; UMC Utrecht Holding B. V.  
 SOURCE: PCT Int. Appl., 55 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

| PATENT NO.  | KIND | DATE     | APPLICATION NO.  | DATE        |
|---|------|----------|--|-------------|
| WO 2005083123   | A1   | 20050909 | WO 2005-NL155  | 20050302    |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |      |          |  |             |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG  |      |          |  |             |
| EP 1571225  | A1   | 20050907 | EP 2004-75686  | 20040302    |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK   |      |          |  |             |
| CA 2558604  | A1   | 20050909 | CA 2005-2558604  | 20050302    |
| EP 1725679  | A1   | 20061129 | EP 2005-710924   | 20050302    |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR   |      |          |  |             |
| US 20070077578  | A1   | 20070405 | US 2006-514345   | 20060831    |
| US 20090098563  | A1   | 20090416 | US 2008-284203   | 20080919    |
| PRIORITY APPLN. INFO.:  |      |          | EP 2004-75686  | A 20040302  |
|   |      |          | US 2004-549450P  | P 20040302  |
|   |      |          | EP 2005-710924   | A 20050302  |
|   |      |          | WO 2005-NL155  | W 20050302  |
|   |      |          | US 2006-514345   | B1 20060831 |
| AB  |      |          | This invention provides methods of detecting CD133 antigen (AC133) |             |

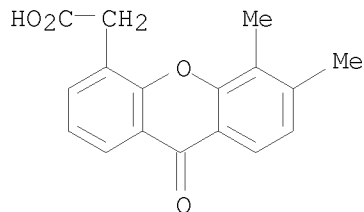
expression level and use as a biomarker for human cancer diagnosis and therapy monitor. Blood anal. including number of circulating endothelial cells and expression levels of human genes AC133 (CD133), EST032 and U1A evaluated by NASBA anal., were determined prior to and during chemotherapy using drugs such as angiostatin or PrimMed01, gemcitabine, and cisplatin, for a wide range of human tumor types. A use of a nucleic acid mol. comprising at least part of a sequence of AC133 or an analog thereof for monitoring a treatment of an individual suffering from a disease is also provided, as well as a diagnostic kit comprising such nucleic acid mol.

IT 95058-81-4, Gemcitabine 117570-53-3, DMXAA  
 RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (methods of detecting CD133 antigen (AC133) expression level and use as biomarker for human cancer diagnosis and therapy monitor)  
 RN 95058-81-4 CAPLUS  
 CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



RN 117570-53-3 CAPLUS  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 4 OF 13 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2005:975665 CAPLUS

DOCUMENT NUMBER: 143:264929

TITLE: Methods for detecting AC133 antigen mRNA for diagnosis and treatment of cancer and other diseases

INVENTOR(S): Penning, Maarten Tjerk; Beerepoot, Laurens Victor; Van Den Broek, Sebastiaan Johannes Jacobus; Mehra, Niven; Voest, Emile Eugene

PATENT ASSIGNEE(S): Primagen Holding B.V., Neth.; UMC Utrecht Holding B.V.  
 SOURCE: Eur. Pat. Appl., 28 pp.

CODEN: EPXXDW

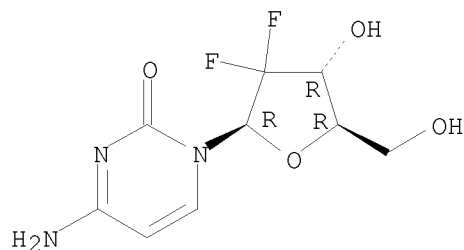
DOCUMENT TYPE: Patent

LANGUAGE: English

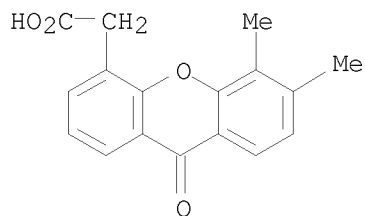
FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO.  | KIND  | DATE     | APPLICATION NO. | DATE       |
|---|---|----------|-----------------|------------|
| EP 1571225  | A1  | 20050907 | EP 2004-75686   | 20040302   |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK   |   |          |                 |            |
| CA 2558604  | A1  | 20050909 | CA 2005-2558604 | 20050302   |
| WO 2005083123   | A1  | 20050909 | WO 2005-NL155   | 20050302   |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |   |          |                 |            |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG  |   |          |                 |            |
| EP 1725679  | A1  | 20061129 | EP 2005-710924  | 20050302   |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR   |   |          |                 |            |
| PRIORITY APPLN. INFO.:  |   |          | EP 2004-75686   | A 20040302 |
|   |   |          | US 2004-549450P | P 20040302 |
|   |   |          | WO 2005-NL155   | W 20050302 |
| AB  | The invention provides methods for detecting AC133 antigen mRNA for diagnosis and treatment of cancer and other diseases. AC133 antigen mRNA may be quantitated by PCR, RT-PCR, NASBA, SDA, TMA, bDNA or rolling circle amplification. Diseases include cancer and heart disease, high blood pressure, ischemia, stroke, psoriasis, Crohn's disease, rheumatoid arthritis, endometriosis, atherosclerosis, obesity, diabetes mellitus, diabetic retinopathy, macular degeneration, Alzheimer's disease, Peutz Jegher's syndrome, multiple sclerosis, systemic lupus erythematosus, Wegener's granulomatosis, vasculitis, sickle cell disease, thalassemia and angina. |          |                 |            |
| IT  | 95058-81-4, Gemcitabine 117570-53-3<br>RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)<br>(methods for detecting AC133 antigen mRNA for diagnosis and treatment of cancer and other diseases)   |          |                 |            |
| RN  | 95058-81-4 CAPLUS   |          |                 |            |
| CN  | Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)  |          |                 |            |
| Absolute stereochemistry. Rotation (+).   |   |          |                 |            |



RN 117570-53-3 CAPLUS  
CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 5 OF 13 CAPLUS COPYRIGHT 2009 ACS on STN  
 ACCESSION NUMBER: 2003:202462 CAPLUS  
 DOCUMENT NUMBER: 138:226761  
 TITLE: Synergistic anticancer combinations containing 5,6-dimethylxanthenone-4-acetic acid  
 INVENTOR(S): Wilson, William Robert; Siim, Bronwyn Gae  
 PATENT ASSIGNEE(S): Cancer Research Technology Limited, UK  
 SOURCE: PCT Int. Appl., 31 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

| PATENT NO.  | KIND | DATE     | APPLICATION NO.  | DATE     |
|---|------|----------|------------------|----------|
| WO 2003020259   | A2   | 20030313 | WO 2002-GB4025   | 20020903 |
| WO 2003020259   | A3   | 20030417 |                  |          |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW |      |          |                  |          |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG  |      |          |                  |          |
| CA 2458459  | A1   | 20030313 | CA 2002-2458459  | 20020903 |
| AU 2002324143   | A1   | 20030318 | AU 2002-324143   | 20020903 |
| AU 2002324143   | B2   | 20070913 |                  |          |
| EP 1423105  | A2   | 20040602 | EP 2002-758562   | 20020903 |
| EP 1423105  | B1   | 20081203 |                  |          |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK   |      |          |                  |          |
| BR 2002012258   | A    | 20041019 | BR 2002-12258    | 20020903 |
| JP 2005509599   | T    | 20050414 | JP 2003-524567   | 20020903 |
| CN 1708296  | A    | 20051214 | CN 2002-817257   | 20020903 |
| NZ 531045   | A    | 20060831 | NZ 2002-531045   | 20020903 |
| EP 1759694  | A2   | 20070307 | EP 2006-77049    | 20020903 |
| EP 1759694  | A3   | 20090218 |                  |          |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE, SK, TR, AL, LT, LV, MK, RO, SI   |      |          |                  |          |
| NZ 546573   | A    | 20070531 | NZ 2002-546573   | 20020903 |
| CN 1994287  | A    | 20070711 | CN 2006-10151393 | 20020903 |
| NZ 554093   | A    | 20080731 | NZ 2002-554093   | 20020903 |
| AT 415963   | T    | 20081215 | AT 2002-758562   | 20020903 |
| NO 2004000591   | A    | 20040430 | NO 2004-591      | 20040210 |

|                        |    |          |                |             |
|------------------------|----|----------|----------------|-------------|
| ZA 2004001078          | A  | 20050415 | ZA 2004-1078   | 20040210    |
| US 20040204480         | A1 | 20041014 | US 2004-790943 | 20040302    |
| MX 2004002004          | A  | 20050217 | MX 2004-2004   | 20040302    |
| IN 2004CN00684         | A  | 20060113 | IN 2004-CN684  | 20040402    |
| US 20070060637         | A1 | 20070315 | US 2006-592678 | 20061103    |
| AU 2007202083          | A1 | 20070531 | AU 2007-202083 | 20070509    |
| US 20080070847         | A1 | 20080320 | US 2007-830650 | 20070730    |
| US 20080070848         | A1 | 20080320 | US 2007-830659 | 20070730    |
| US 20080070886         | A1 | 20080320 | US 2007-830668 | 20070730    |
| US 20080070849         | A1 | 20080320 | US 2007-830677 | 20070730    |
| PRIORITY APPLN. INFO.: |    |          | GB 2001-21285  | A 20010903  |
|                        |    |          | AU 2002-324143 | A3 20020903 |
|                        |    |          | CN 2002-817257 | A3 20020903 |
|                        |    |          | EP 2002-758562 | A3 20020903 |
|                        |    |          | WO 2002-GB4025 | W 20020903  |
|                        |    |          | US 2004-790943 | A1 20040302 |

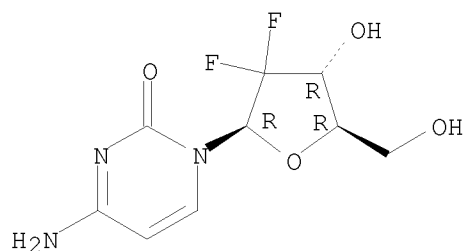
AB The present invention relates to synergistic combinations of the 5,6-dimethylxanthenone-4-acetic acid (DMXAA) and a compound selected from platinum compds., Vinca alkaloids, alkylating agents, anthracyclines, topoisomerase I inhibitors, antimetabolites and topoisomerase II inhibitors, which have antitumor activity. More particularly, the invention is concerned with the use of such combinations in the treatment of cancer and pharmaceutical compds. containing the combinations. The antitumor activity and host toxicity of DMXAA/cytotoxic drug combinations was assessed by varying the dose of chemotherapeutic drug up to the toxicity limit, with co-administration of a fixed DMXAA dose (80  $\mu$ mol/kg, ca. 80% of MTD), and evaluating subsequent tumor growth delay. Of the 7 drugs investigated, 4 (doxorubicin, 5-fluorouracil, cyclophosphamide and cisplatin) had appreciable activity against this tumor as indicated by dose-response relationships providing significant slopes by linear regression, and highly significant growth delays of 10 days at their MTDs.

IT 95058-81-4, Gemcitabine  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (synergistic anticancer combinations)

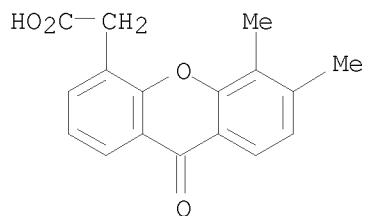
RN 95058-81-4 CAPLUS

CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



IT 117570-53-3, 5,6-Dimethylxanthenone-4-acetic acid  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (synergistic anticancer combinations containing dimethylxanthenoneacetic acid)  
 RN 117570-53-3 CAPLUS  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 13 USPATFULL on STN  
 ACCESSION NUMBER: 2008:80755 USPATFULL  
 TITLE: ANTI-CANCER COMBINATIONS  
 INVENTOR(S): Wilson, William R., Waiuku, NEW ZEALAND  
 Siim, Bronwyn G., Auckland, NEW ZEALAND  
 PATENT ASSIGNEE(S): CANCER RESEARCH TECHNOLOGY LIMITED, London, UNITED KINGDOM (non-U.S. corporation)

|                       | NUMBER  | KIND | DATE          |
|-----------------------|---|------|---------------|
| PATENT INFORMATION:   | US 20080070886  | A1   | 20080320      |
| APPLICATION INFO.:    | US 2007-830668  | A1   | 20070730 (11) |
| RELATED APPLN. INFO.: | Continuation of Ser. No. US 2004-790943, filed on 2 Mar 2004, PENDING |      |               |

|                       | NUMBER   | DATE     |
|-----------------------|--|----------|
| PRIORITY INFORMATION: | WO 2002-GB4025   | 20020903 |
|                       | GB 2001-21285  | 20010903 |
| DOCUMENT TYPE:        | Utility  |          |
| FILE SEGMENT:         | APPLICATION  |          |
| LEGAL REPRESENTATIVE: | JAECKLE FLEISCHMANN & MUGEL, LLP, 190 Linden Oaks, ROCHESTER, NY, 14625-2812, US |          |
| NUMBER OF CLAIMS:     | 23   |          |
| EXEMPLARY CLAIM:      | 1  |          |
| NUMBER OF DRAWINGS:   | 2 Drawing Page(s)  |          |
| LINE COUNT:           | 1275   |          |

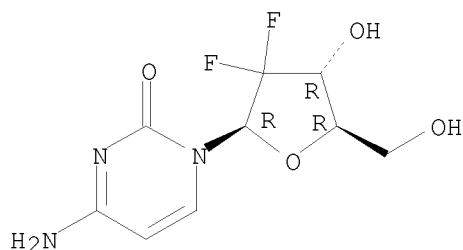
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates to synergistic combinations of the compound 5,6-dimethylxanthenone-4-acetic acid (DMXAA) and a compound selected from platinum compounds, vinca alkaloids, alkylating agents, anthracyclines, topoisomerase I inhibitors, antimetabolites and topoisomerase II inhibitors, which have anti-tumour activity. Preferably, the present invention relates to synergistic combinations of the compound 5,6-dimethylxanthenone-4-acetic acid (DMXAA) and a compound selected from carboplatin, gemcitabine, cisplatin, 5-fluorouracil, cyclophosphamide, etoposide, vincristine, doxorubicin and irinotecan. More particularly, the invention is concerned with the use of such combinations in the treatment of cancer and pharmaceutical compositions containing such combinations. The invention further provides for methods of preparing the combinations of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 95058-81-4, Gemcitabine  
 (synergistic anticancer combinations)  
 RN 95058-81-4 USPATFULL  
 CN Cytidine, 2'-deoxy-2',2'-difluoro- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



=> d his

(FILE 'HOME' ENTERED AT 11:33:44 ON 17 APR 2009)

FILE 'REGISTRY' ENTERED AT 11:34:01 ON 17 APR 2009

E "DMXAA"/CN 25

L1 1 S E3

E "GEMCITABINE"/CN 25

L2 1 S E3

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 11:34:54 ON 17 APR 2009

L3 370 S L1

L4 5732 S L2

L5 13 S L3 AND L4

=> s l3 and antimetabolite

L6 8 L3 AND ANTIMETABOLITE

=> d l6 1-8 ibib, abs, hitstr

L6 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2004:284727 CAPLUS

DOCUMENT NUMBER: 142:85467

TITLE: The Cancer Research UK experience of pre-clinical toxicology studies to support early clinical trials with novel cancer therapies

AUTHOR(S): Newell, D. R.; Silvester, J.; McDowell, C.; Burtles, S. S.

CORPORATE SOURCE: Cancer Research UK, Drug Development Office, London, WC2A 3PX, UK

SOURCE: European Journal of Cancer (2004), 40(6), 899-906  
CODEN: EJCAEL; ISSN: 0959-8049

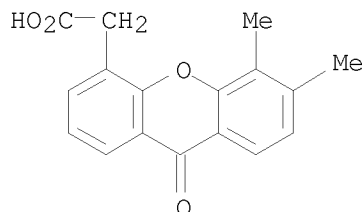
PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. Pre-clin. toxicol. studies in rodents and Phase I clin. trial data are summarized for 14 novel anticancer therapies. With only one exception, an antifolate antimetabolite, rodent toxicol. predicted a safe Phase I trial starting dose and the majority of the dose limiting toxicities, in particular haematol. toxicity. For targeted agents with well-defined pharmacodynamic markers, illustrated in the current study by 3 anti-endocrine drugs and one resistance modifier, the definition of a maximum tolerated dose can be avoided. Together with earlier data, the current study confirms that pre-clin. toxicol. studies in a non-rodent species are not routinely needed for the safe conduct of early clin. trials with new cancer chemotherapies.

IT 117570-53-3, DMXAA  
 RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (cancer research UK experience of pre-clin. toxicol. studies to support  
 early clin. trials with novel cancer therapies)  
 RN 117570-53-3 CAPLUS  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS  
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 8 USPATFULL on STN  
 ACCESSION NUMBER: 2008:80755 USPATFULL  
 TITLE: ANTI-CANCER COMBINATIONS  
 INVENTOR(S): Wilson, William R., Waiuku, NEW ZEALAND  
 Siim, Bronwyn G., Auckland, NEW ZEALAND  
 PATENT ASSIGNEE(S): CANCER RESEARCH TECHNOLOGY LIMITED, London, UNITED  
 KINGDOM (non-U.S. corporation)

|                       | NUMBER  | KIND | DATE          |
|-----------------------|---|------|---------------|
| PATENT INFORMATION:   | US 20080070886  | A1   | 20080320      |
| APPLICATION INFO.:    | US 2007-830668  | A1   | 20070730 (11) |
| RELATED APPLN. INFO.: | Continuation of Ser. No. US 2004-790943, filed on 2 Mar 2004, PENDING |      |               |

|                       | NUMBER   | DATE     |
|-----------------------|--|----------|
| PRIORITY INFORMATION: | WO 2002-GB4025   | 20020903 |
|                       | GB 2001-21285  | 20010903 |
| DOCUMENT TYPE:        | Utility  |          |
| FILE SEGMENT:         | APPLICATION  |          |
| LEGAL REPRESENTATIVE: | JAECKLE FLEISCHMANN & MUGEL, LLP, 190 Linden Oaks, ROCHESTER, NY, 14625-2812, US |          |
| NUMBER OF CLAIMS:     | 23   |          |
| EXEMPLARY CLAIM:      | 1  |          |
| NUMBER OF DRAWINGS:   | 2 Drawing Page(s)  |          |
| LINE COUNT:           | 1275   |          |

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates to synergistic combinations of the compound 5,6-dimethylxanthene-4-acetic acid (DMXAA) and a compound selected from platinum compounds, vinca alkaloids, alkylating agents, anthracyclines, topoisomerase I inhibitors, antimetabolites and topoisomerase II inhibitors, which have anti-tumour activity. Preferably, the present invention relates to synergistic combinations of the compound 5,6-dimethylxanthene-4-acetic acid (DMXAA) and a compound selected from carboplatin, gemcitabine, cisplatin, 5-fluorouracil, cyclophosphamide, etoposide, vincristine, doxorubicin and irinotecan. More particularly, the invention is concerned with the use of such combinations in the treatment of cancer and pharmaceutical compositions containing such combinations. The invention further provides for methods



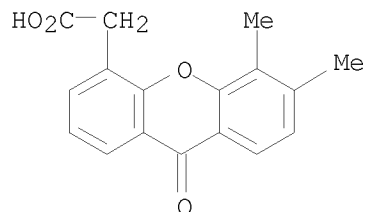
of preparing the combinations of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 117570-53-3, 5,6-Dimethylxanthenone-4-acetic acid  
(synergistic anticancer combinations containing dimethylxanthenoneacetic acid)

RN 117570-53-3 USPATFULL

CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



=> d his

(FILE 'HOME' ENTERED AT 11:33:44 ON 17 APR 2009)

FILE 'REGISTRY' ENTERED AT 11:34:01 ON 17 APR 2009

E "DMXAA"/CN 25

L1 1 S E3

E "GEMCITABINE"/CN 25

L2 1 S E3

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 11:34:54 ON 17 APR 2009

L3 370 S L1

L4 5732 S L2

L5 13 S L3 AND L4

L6 8 S L3 AND ANTIMETABOLITE

=> s l3 and (?potentiat? or ?enhanc? or ?increas?)

L7 228 L3 AND (?POTENTIAT? OR ?ENHANC? OR ?INCREAS?)

=> s l7 and (?cancer? or ?tumor? or ?tumour? or ?neoplasm?)

L8 224 L7 AND (?CANCER? OR ?TUMOR? OR ?TUMOUR? OR ?NEOPLASM?)

=> s l8 and (pd<20020903 or prd<20020903)

'20020903' NOT A VALID FIELD CODE

2 FILES SEARCHED...

3 FILES SEARCHED...

L9 132 L8 AND (PD<20020903 OR PRD<20020903)

=> dup rem l9

PROCESSING COMPLETED FOR L9

L10 83 DUP REM L9 (49 DUPLICATES REMOVED)

=> s l10 and antimetabolite

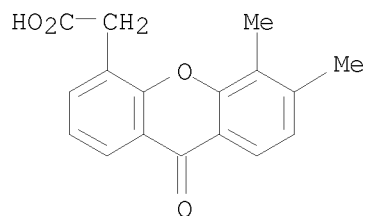
L11 6 L10 AND ANTIMETABOLITE

=> d l6 l-11 ibib, abs, hitstr

L6 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2004:284727 CAPLUS

DOCUMENT NUMBER: 142:85467  
 TITLE: The Cancer Research UK experience of pre-clinical toxicology studies to support early clinical trials with novel cancer therapies  
 AUTHOR(S): Newell, D. R.; Silvester, J.; McDowell, C.; Burtles, S. S.  
 CORPORATE SOURCE: Cancer Research UK, Drug Development Office, London, WC2A 3PX, UK  
 SOURCE: European Journal of Cancer (2004), 40(6), 899-906  
 CODEN: EJCAEL; ISSN: 0959-8049  
 PUBLISHER: Elsevier Science Ltd.  
 DOCUMENT TYPE: Journal; General Review  
 LANGUAGE: English  
 AB A review. Pre-clin. toxicol. studies in rodents and Phase I clin. trial data are summarized for 14 novel anticancer therapies. With only one exception, an antifolate antimetabolite, rodent toxicol. predicted a safe Phase I trial starting dose and the majority of the dose limiting toxicities, in particular haematol. toxicity. For targeted agents with well-defined pharmacodynamic markers, illustrated in the current study by 3 anti-endocrine drugs and one resistance modifier, the definition of a maximum tolerated dose can be avoided. Together with earlier data, the current study confirms that pre-clin. toxicol. studies in a non-rodent species are not routinely needed for the safe conduct of early clin. trials with new cancer chemotherapies.  
 IT 117570-53-3, DMXAA  
 RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (cancer research UK experience of pre-clin. toxicol. studies to support early clin. trials with novel cancer therapies)  
 RN 117570-53-3 CAPLUS  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 8 USPATFULL on STN

ACCESSION NUMBER: 2008:80755 USPATFULL  
 TITLE: ANTI-CANCER COMBINATIONS  
 INVENTOR(S): Wilson, William R., Waiuku, NEW ZEALAND  
 Siim, Bronwyn G., Auckland, NEW ZEALAND  
 PATENT ASSIGNEE(S): CANCER RESEARCH TECHNOLOGY LIMITED, London, UNITED KINGDOM (non-U.S. corporation)

|                       | NUMBER  | KIND | DATE          |
|-----------------------|---|------|---------------|
| PATENT INFORMATION:   | US 20080070886  | A1   | 20080320      |
| APPLICATION INFO.:    | US 2007-830668  | A1   | 20070730 (11) |
| RELATED APPLN. INFO.: | Continuation of Ser. No. US 2004-790943, filed on 2 Mar 2004, PENDING |      |               |

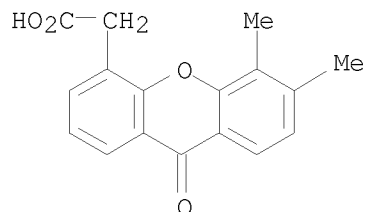
| NUMBER | DATE |
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 PRIORITY INFORMATION: WO 2002-GB4025 20020903  
 GB 2001-21285 20010903  
 DOCUMENT TYPE: Utility  
 FILE SEGMENT: APPLICATION  
 LEGAL REPRESENTATIVE: JAECKLE FLEISCHMANN & MUGEL, LLP, 190 Linden Oaks,  
 ROCHESTER, NY, 14625-2812, US  
 NUMBER OF CLAIMS: 23  
 EXEMPLARY CLAIM: 1  
 NUMBER OF DRAWINGS: 2 Drawing Page(s)  
 LINE COUNT: 1275  
 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates to synergistic combinations of the compound 5,6-dimethylxanthenone-4-acetic acid (DMXAA) and a compound selected from platinum compounds, vinca alkaloids, alkylating agents, anthracyclines, topoisomerase I inhibitors, antimetabolites and topoisomerase II inhibitors, which have anti-tumour activity. Preferably, the present invention relates to synergistic combinations of the compound 5,6-dimethylxanthenone-4-acetic acid (DMXAA) and a compound selected from carboplatin, gemcitabine, cisplatin, 5-fluorouracil, cyclophosphamide, etoposide, vincristine, doxorubicin and irinotecan. More particularly, the invention is concerned with the use of such combinations in the treatment of cancer and pharmaceutical compositions containing such combinations. The invention further provides for methods of preparing the combinations of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 117570-53-3, 5,6-Dimethylxanthenone-4-acetic acid  
 (synergistic anticancer combinations containing dimethylxanthenoneacetic acid)  
 RN 117570-53-3 USPATFULL  
 CN 9H-Xanthene-4-acetic acid, 5,6-dimethyl-9-oxo- (CA INDEX NAME)



=> d his

(FILE 'HOME' ENTERED AT 11:33:44 ON 17 APR 2009)

FILE 'REGISTRY' ENTERED AT 11:34:01 ON 17 APR 2009  
 E "DMXAA"/CN 25

L1 1 S E3  
 E "GEMCITABINE"/CN 25  
 L2 1 S E3

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 11:34:54 ON 17 APR 2009

L3 370 S L1  
 L4 5732 S L2  
 L5 13 S L3 AND L4  
 L6 8 S L3 AND ANTIMETABOLITE

L7 228 S L3 AND (?POTENTIAT? OR ?ENHANC? OR ?INCREAS?)  
L8 224 S L7 AND (?CANCER? OR ?TUMOR? OR ?TUMOUR? OR ?NEOPLASM?)  
L9 132 S L8 AND (PD<20020903 OR PRD<20020903)  
L10 83 DUP REM L9 (49 DUPLICATES REMOVED)  
L11 6 S L10 AND ANTIMETABOLITE

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---Logging off of STN---

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Executing the logoff script...

=> LOG Y

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|--|------------|---------|
| COST IN U.S. DOLLARS                       | SINCE FILE | TOTAL   |
|  | ENTRY      | SESSION |
| FULL ESTIMATED COST                        | 103.87     | 119.85  |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL   |
|  | ENTRY      | SESSION |
| CA SUBSCRIBER PRICE                        | -5.74      | -5.74   |

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